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Neutrokin- $\alpha$ 

1 AAATTCAAGGATAACTCTCTGAGGGGTGAGCCAAGCCTGCCATGTAGTGCACGCAGGAC 60  
 61 ATCAACAAACACAGATAACAGGAAATGATCCATTCCCTGTGGTCACTTATTCTAAAGGCC 120  
 121 CCAACCTTCAAAGTTCAAGTAGTGATATGGATGACTCCACAGAAAGGGAGCAGTCACGCC 180  
 1 M D D S T E R E Q S R L 12  
 181 TTACTTCTTGCCCTTAAGAAAAGAGAAAGAAATGAACTGAAGGAGTGTGTTTCCATCCTCC 240  
 13 T S C L K K R E E M K L K E C V S I L P 32  
 CD-I  
 241 CACGGAAGGAAAGCCCTCTGTCCGATCCTCCAAAGACGGAAGCTGCTGGTGCAACCT 300  
 33 R K E S P S V R S S K D G K L L A A T L 52  
 CD-I  
 301 TGCTGCTGGCACTGCTGTCTTGCTGCCTCACGGTGGTGTCTTTCTACCAAGTGCGCGCCC 360  
 53 L L A L L S C C L T V V S F Y Q V A A L 72  
 361 TGCAAGGGGACCTGGCCAGCCTCCGGGCAGAGCTGCAGGGCCACCACGCGGAGAAGCTGC 420  
 73 Q G D L A S L R A E L Q G H H A E K L P 92  
 CD-II  
 421 CAGCAGGAGCAGGAGCCCCAAGGCCGGCCTGGAGGAAGCTCCAGCTGTACCGCGGGAC 480  
 93 A G A G A P K A G L E E A P A V T A G L 112  
 CD-III  
 #  
 481 TGAAATCTTTGAACCAACAGCTCCAGGAGAAGGCAACTCCAGTCAGAACAGCAGAAAT 540  
 113 K I F E P P A P G E G N S S Q N S R N K 132  
 541 AGCGTGCCGTTCAAGGTCCAGAAGAAACAGTCAAGACTGCTTGCAACTGATTGCAG 600  
 133 R A V Q G P E E T V T Q D C L Q L I A D 152  
 CD-IV

FIG.1A

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Neutrokin- $\alpha$ 

601	ACAGTGAACACCAACTATACAAAAAGGATCTTACACATTGTTCATGGCTTCTCAGCT	660
153	S E T P T I Q K G S Y T F V P W L L S F	172
	CD-V	
661	TTAAAAGGGGAAGTCCCTAGAAAGAAAAGAGAATAAAATATTGGTCAAAGAACTGGTT	720
173	K R G S A L E E K E N K I L V K E T G Y	192
	CD-V	CD-VI
721	ACTTTTTTATATATGGTCAGGTTTTATATACTGATAAGACCTACGCCATGGGACATCTAA	780
193	F F I Y G Q V L Y T D K T Y A M G H L I	212
	CD-VI	CD-VII
781	TTCAGAGGAAGAAGGTCCATGTCTTTGGGGATGAATTGAGTCTGGTGACTTTGTTTCGAT	840
213	Q R K K V H V F G D E L S L V T L F R C	232
	CD-VII	CD-VIII
	#	
841	GTATTCAAAATATGCCTGAAACACTACCCAATAATTCCTGCTATTAGCTGGCATTGCAA	900
233	I Q N M P E T L P N N S C Y S A G I A K	252
	CD-VIII	CD-IX
901	AACTGGAAGAAGGAGATGAACTCCAACCTGCAATACCAAGAGAAAAATGCACAAATATCAC	960
253	L E E G D E L Q L A I P R E N A Q I S L	272
	CD-X	
961	TGGATGGAGATGTCACATTTTTTGGTGCAATTGAAACTGCTGTGACCTACTTACACCATGT	1020
273	D G D V T F F G A L K L L	285
	CD-XI	
1021	CTGTAGCTATTTTCTCCCTTTCTCTGTACCTCTAAGAAGAAAGAAATCTAACTGAAAATA	1080
1081	CCAAAAA	1100

FIG.1B

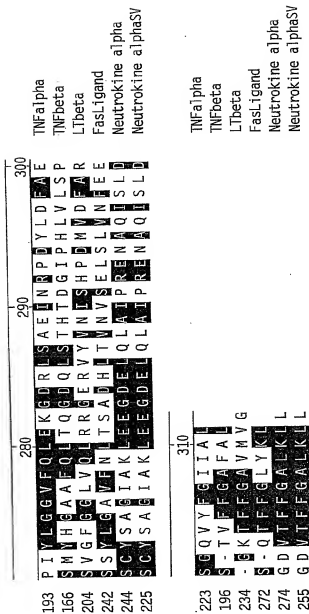
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		10	20	30	
1	STESMIRDVEL				TNFalpha
1				- - - - - AEEA	TNFbeta
1				- - - - - TPPERL	Lbeta
1	GA			- - - - -	FasLigand
1	QQPFNYYPQIYW			- VDSASSPWAPPGTV	Neutrokine alpha
1	DDSTEREQSRL			TSCCLKREEMKLKECVSI	Neutrokine alphaSV
1	DDSTEREQSRL			TSCCLKREEMKLKECVSI	
		40	50	60	
17	LPKKTGGPQ				TNFalpha
8	F				TNFbeta
4					Lbeta
30	LPCTISVPRP				FasLigand
31	LPRESPSVRS				Neutrokine alpha
31	LPRESPSVRS				Neutrokine alphaSV
		70	80	90	
30					TNFalpha
9					TNFbeta
12					Lbeta
60	PPPLPPLP				FasLigand
58	SCCLTVVSF				Neutrokine alpha
58	SCCLTVVSF				Neutrokine alphaSV

FIG.2A

FIG. 2B

FIG. 2C



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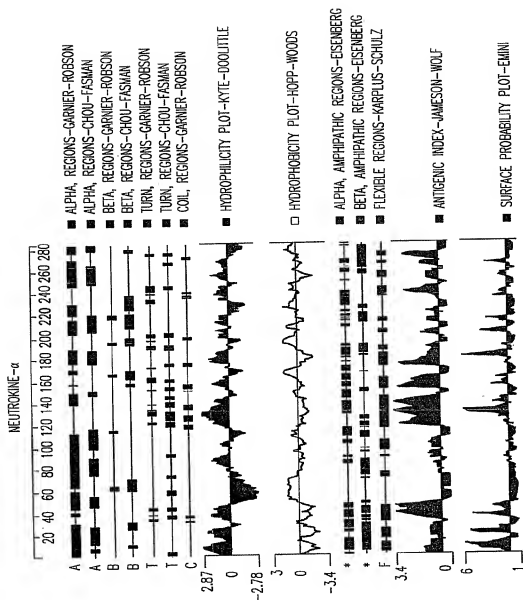


FIG.3

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	1				50
HSOAD55R	.....A	GGNTAACTCT	CCTGAGGGGT	GAGCCAAGCC	CTGCCATGTA
HNEDU15X	...AAATTCA	GGATAACTCT	CCTGAGGGGT	GAGCCAAGCC	CTGCCATGTA
HSLAH84R	.AATTGGGCA	NAGNAACTG	GTTACTTTTT	TATATATGGT	CAGGTTTAT
HLTBM08R	AATTCGGCAC	GAGCAAGGCC	GGCCTGGAGG	AAGCTCCAGC	TGTCACCGCG
	51				100
HSOAD55R	GTGCACGCAG	GACATCANCA	A..ACACANN	NNNCAGGAAA	TAATCCATTTC
HNEDU15X	GTGCACGCAG	GACATCAACA	A..ACACAGA	TAACAGGAAA	TGATCCATTTC
HSLAH84R	ATACTGATAA	GACCTACGCC	ATGGGACATC	TAGTTCAGAG	GAAGAAGGTC
HLTBM08R	GGACTGAAAA	TCTTTGAACC	ACCAGCTCCA	GGAGAAGGCA	ACTCCAGTCA
	101				150
HSOAD55R	CCTGTGGTCA	CTTATTCTAA	AGGCCCAAC	CTTCAAAGTT	CAAGTAGTGA
HNEDU15X	CCTGTGGTCA	CTTATTCTAA	AGGCCCAAC	CTTCAAAGTT	CAAGTAGTGA
HSLAH84R	CATGTCTTTG	GGGATGAATT	GAGTCTGGTG	ACTTTGTTTC	GATGTATTCA
HLTBM08R	GAACAGCAGA	AATAAGCGTG	CCGTTACGGG	TCCAGAAGAA	ACAGTCACTC
	151				200
HSOAD55R	TATGGATGAC	TCCACAGAAA	GGGAGCAGTC	ACGCCCTTACT	TCTTGCCCTTA
HNEDU15X	TATGGATGAC	TCCACAGAAA	GGGAGCAGTC	ACGCCCTTACT	TCTTGCCCTTA
HSLAH84R	AAATATGCCT	GAACACTAC	CCAATAATTC	CTGCTATTCA	GCTGGCATTG
HLTBM08R	AAGACTGCCT	GCAACTGNTT	GCAGACAGTG	AAACACCAAC	TATACAAAAA
	201				250
HSOAD55R	AGAAAAAGAGA	AGAAATGAAA	CTGNAAGGAG	TGTGTTTCCA	TCCTCCCACG
HNEDU15X	AGAAAAAGAGA	AGAAATGAAA	CT..GAAGGAG	TGTGTTTCCA	TCCTCCCACG
HSLAH84R	CAAAACTGGN	AGGAAGGA..	...GATGAAC	TCCAAC TTG	AATACCAGGG
HLTBM08R	GGCTCCCTTC	TGNTGCCACA	TTTGGGCCAA	GGATGGAGA	GATTTCTTCG
	251				300
HSOAD55R	GAAGGAAAGC	CCCTCTNTCC	GATCCTCCAA	AGACGGAAAG	CTGCTGGCTG
HNEDU15X	GAAGGAAAGC	CCCTCTGTCC	GATCCTCCAA	AGACGGAAAG	CTGCTGGCTG
HSLAH84R	GAAAATGCAC	AATTATCACT	GGGATGGAGA	TGTTACACT	TTTTGGGTGC
HLTBM08R	TCTGGAACA	TTTTGCCAAA	CTCTTCAGAT	ACTCTTTNCT	CTCTGGGAAT
	301				350
HSOAD55R	CAACCTTGNT	GNTGGCATTG	TGTCTTGCT	GNCTCAAGGT	GGTGTNTTT.
HNEDU15X	CAACCTTGCT	GCTGGCATTG	CTGTCTTGCT	GCCTCACGGT	GGTGTCTTTT
HSLAH84R	CATTGAAACT	GCTGTGACCT	NCTTACANCA	NGTCTGTNT	GCTATTTTTC
HLTBM08R	CAAAGGAAAA	TCTCTACTTA	GATTNACACA	TTTGTTCCTCA	TGGGTNTCTT
	351				400
HSOAD55R	.....	.....	.....	.....	.....
HNEDU15X	TACCAGGTGG	CCGCCCTGCA	AGGGGACCTG	GCCAGCCTCC	GGGACAGAGCT
HSLAH84R	CTNCTNTTTC	TNTGGTAACC	TCTTAGGAAG	GAAGGATTCT	TAACTGGGAA
HLTBM08R	AAGTTTTAAA	AGGGGAGTGC	CCTTAGGAGG	AAAAGGGGAT	AAATATTGGC

FIG.4A



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	401		450
HSOAD55R	.....	.....	.....
HNEDU15X	GCAGGGCCAC	CACGCGGAGA	AGCTGCCAGC
HSLAH84R	ATAACCCAAA	AAAANNTTAA	ANGGGTANGN
HLTBM08R	CAAGGNACTG	GTTANTTTNT	AAATATGGTC
			AGGTTTNTAT
			ANCTGGTAGG
	451		500
HSOAD55R	.....	.....	.....
HNEDU15X	CCGGCCTGGA	GGAAGCTCCA	GCTGTCCACG
HSLAH84R	CNNNGNNGNT	TTTNGGNNTA	TNTTNTNNTN
HLTBM08R	CCTGCCCATG	GGCATTNATT	CANGGNGAGG
			NCNNTCTTTT
			GGGNTGA...
	501		550
HSOAD55R	.....	.....	.....
HNEDU15X	CCACCAGCTC	CAGGAGAAGG	CAACTCCAGT
HSLAH84R	CNANGGGGGN	TTTTT.....	.....
HLTBM08R	.....	.....	.....
			.....
			.....
	551		600
HSOAD55R	.....	.....	.....
HNEDU15X	TGCCGTTTCA	GGTCCAGAAG	AAACAGTCAC
HSLAH84R	.....	.....	TCAAGACTGC
HLTBM08R	.....	.....	TTGCAACTGA
			.....
			.....
	601		650
HSOAD55R	.....	.....	.....
HNEDU15X	TTGCAGACAG	TGAAACACCA	ACTATACAAA
HSLAH84R	.....	.....	AAGGATCTTA
HLTBM08R	.....	.....	CACATTTTGT
			.....
			.....
	651		700
HSOAD55R	.....	.....	.....
HNEDU15X	CCATGGCTTC	TCAGCTTTAA	AAGGGGAAGT
HSLAH84R	.....	.....	GCCCTAGAAG
HLTBM08R	.....	.....	AAAAAGAGAA
			.....
			.....
	701		750
HSOAD55R	.....	.....	.....
HNEDU15X	TAAAAATATTG	GTCAAAGAAA	CTGGTTACTT
HSLAH84R	.....	.....	TTTTATATAT
HLTBM08R	.....	.....	GGTCAGGTTT
			.....
			.....
	751		800
HSOAD55R	.....	.....	.....
HNEDU15X	TATATACTGA	TAAGACCTAC	GCCATGGGAC
HSLAH84R	.....	.....	ATCTAATTCA
HLTBM08R	.....	.....	GAGGAAGAAG
			.....
			.....

FIG.4B

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	801		850
HSOAD55R	.....	.....	.....
HNEDU15X	GTCCATGTCT	TTGGGGATGA	ATTGAGTCTG GTGACTTTGT TTCGATGTAT
HSLAH84R	.....	.....	.....
HLTBM08R	.....	.....	.....
	851		900
HSOAD55R	.....	.....	.....
HNEDU15X	TCAAAATATG	CCTGAAACAC	TACCCAATAA TTCCTGCTAT TCAGCTGGCA
HSLAH84R	.....	.....	.....
HLTBM08R	.....	.....	.....
	901		950
HSOAD55R	.....	.....	.....
HNEDU15X	TTGCAAAACT	GGAAGAAGGA	GATGAACTCC AACTTGCAAT ACCAAGAGAA
HSLAH84R	.....	.....	.....
HLTBM08R	.....	.....	.....
	951		1000
HSOAD55R	.....	.....	.....
HNEDU15X	AATGCACAAA	TATCACTGGA	TGGAGATGTC ACATTTTTTG GTGCATTGAA
HSLAH84R	.....	.....	.....
HLTBM08R	.....	.....	.....
	1001		1050
HSOAD55R	.....	.....	.....
HNEDU15X	ACTGCTGTGA	CCTACTTACA	CCATGTCTGT AGCTATTTTC CTCCTTTCT
HSLAH84R	.....	.....	.....
HLTBM08R	.....	.....	.....
	1051		1100
HSOAD55R	.....	.....	.....
HNEDU15X	CTGTACCTCT	AAGAAGAAAG	AATCTAAGTG AAAATACCAA AAAAAAAAAA
HSLAH84R	.....	.....	.....
HLTBM08R	.....	.....	.....
	1101		
HSOAD55R	.....		
HNEDU15X	AAAAAA		
HSLAH84R	.....		
HLTBM08R	.....		

FIG.4C

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Neutrokin- $\alpha$ SV

1	ATGGATGACTCCACAGAAAGGGAGCAGTCACGCCTTACTTCTTGCCTTAAGAAAAGAGAA	60
1	M D D S T E R E Q S R L T S C L K K E	20
61	GAAATGAACTGAAGGAGTGTGTTCCATCCTCCACGGAAGGAAAGCCCTCTGTCCGA	120
21	E M K L K E C V S I L P R K E S P S V R	40
	CD-I	
121	TCCTCCAAAGACGGAAGCTGCTGGCTGCAACCTTGCTGCTGGCACTGCTGCTTGTCTGC	180
41	S S K D G K L L A A T L L L A L L S C C	60
	CD-I	
181	CTCACGGTGGTGTCTTTTACCAGGTGGCCGCCCTGCAAGGGGACCTGGCCAGCCTCCGG	240
61	L T V V S F Y Q V A A L Q G D L A S L R	80
	CD-II	
241	GCAGAGCTGCAGGGCCACCACGCGGAGAAGCTGCCAGGAGCAGGAGCCCCAAGGCC	300
81	A E L Q G H H A E K L P A G A G A P K A	100
	CD-II	
301	GGCCTGGAGGAAGCTCCAGCTGTACCGCGGGACTGAAAATCTTTGAACCACCAGCTCCA	360
101	G L E E A P A V T A G L K I F E P P A P	120
	CD-III	
	#	
361	GGAGAAGGCAACTCCAGTCAGAACAGCAGAAAATAAGCGTGCCTTCAGGGTCCAGAGAAGAA	420
121	G E G N S S Q N S R N K R A V Q G P E E	140
421	ACAGGATCTTACACATTTGTTCATGGCTTCTCAGCTTTAAAGGGGAAGTGCCCTAGAA	480
141	T G S Y T F V P W L L S F K R G S A L E	160
	CD-IV	
481	GAAAAGAGAATAAAATATTGGTCAAAGAACTGGTTACTTTTTATATATGGTCAGGTT	540
161	E K E N K I L V K E T G Y F F I Y G Q V	180
	CD-IV	
541	TTATATACTGATAAGACCTACGCCATGGGACATCTAATTAGAGGAAGAAGGTCCATGTC	600
181	L Y T D K T Y A M G H L I Q R K K V H V	200
	CD-VI	
	CD-VII	

FIG.5A

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Neutrokin- $\alpha$ SV

601 TTTGGGATGAATTGAGTCTGGTGACTTTGTTTCGATGTATTCAAAATATGCCTGAAACA 660  
 201 F G D E L S L V T L F R C I Q N M P E T 220  
 CD-VIII CD-VIII

661 CTACCCAATAATTCCTGCTATTCACTGGCATTGCAAACTGGAAGAAGGAGATGAATC 720  
 221 L P N N S C Y S A G I A K L E E G D E L 240  
 CD-IX CD-X

721 CAACTTGCAATACCAAGAGAAAAATGCACAAATATCACTGGATGGAGATGTCACATTTTT 780  
 241 Q L A I P R E N A Q I S L D G D V T F F 260  
 CD-X CD-XI

781 GGTGCATTGAACTGCTGTGACCTACTTACCATGTCTGTAGCTATTTTCTCCCTTTC 840  
 261 G A L K L L 266  
 CD-XI

841 TCTGTACCTCTAAGAAGAAAGAATCTAACTGAAAATACCAAAAAAAAAAAAAAAAAAAAA 900  
 901 AAA 903

FIG.5B

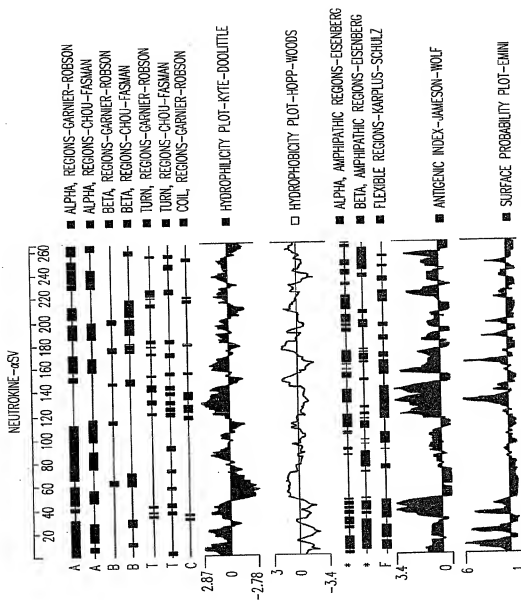


FIG. 6

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Neutrokine-  
Alpha MDDSTEREQSRLLTSCCLKKEEMKLEKCVSILPRKESPSVRS 41

Transmembrane Region

SKDGKLLAATLLALLSCCLTVVSFYQVAALQGDLASLRAE 82

LQGHHA EKLPAGAGAPKAGLEEA PAVTAGLKIFEPAPAGEG 123

↓  
NSSQNSRNKRAVQGPPEETVTQDCQLIADSEIPTIQKGSY 164  
Apr11 HSLHLVPINAISK-DDSDV 134  
TNF KPVAVVAVNQAEGQ----- 102  
LTα KPAALHLIGDPSKQNS----- 76

A' B' C  
FVPM LLS---FKRGSAL EEKENKI L KETGYFFLYGQV L 200  
EVMMPQA-----LRGRGLQAQGYGVRIQDAQV LLYSQV L 170  
-LQMLNRRANALLANGVELRD--NQLVVPSEGLV LLYSQV L 139  
-LLWRANTDRAFLQDGFSLSN--NSLVPTSGLIYFVVSQV V 114

D E  
YT DKT V-----AMGH L IQKKV HVFGDELSLV L LFRCLIQNMP 237  
EQQVTF-----MGQVVSRE-----GRQE L LFRCLIRSM P 201  
FKGQGC P-----STLV L L TISRIVAVSQTKVNLLSAIKSP 176  
ESGKA VSPKATSSPIYLAEEVQLFESSQV PFHVPLLSSQK V 155

FIG.7A-1



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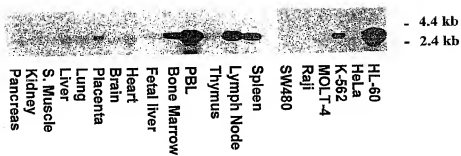


FIG.7B



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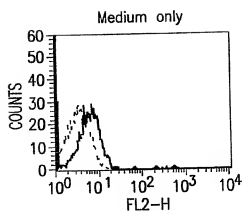


FIG.8A

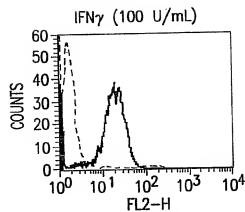


FIG.8B

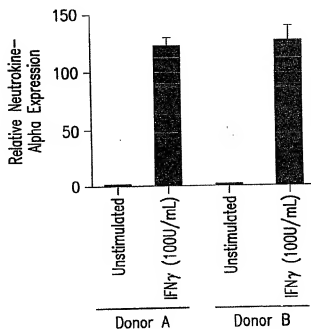


FIG.8C

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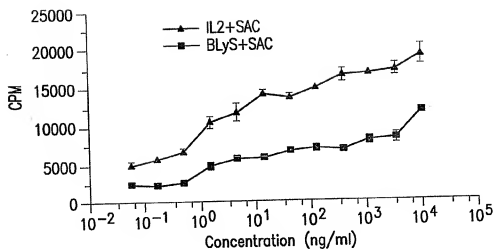


FIG. 9A

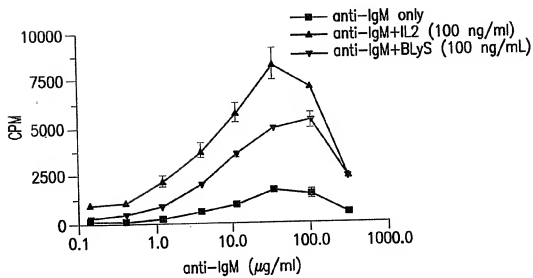


FIG. 9B

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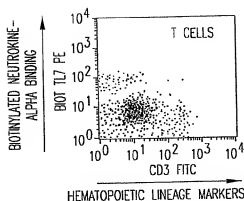


FIG.10A

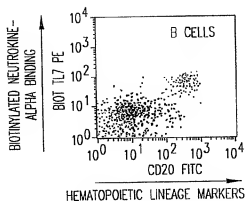


FIG.10B

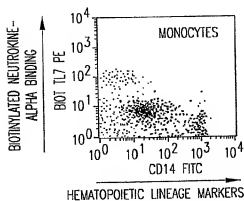


FIG.10C

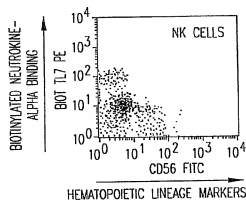


FIG.10D

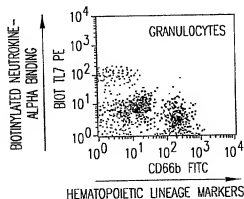


FIG.10E

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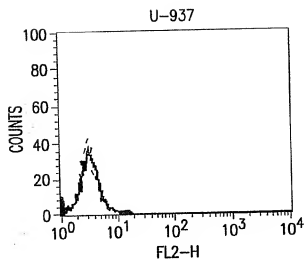


FIG.10F

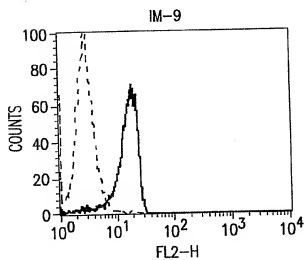


FIG.10G

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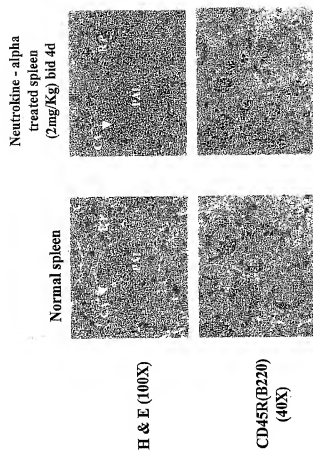


FIG.11A

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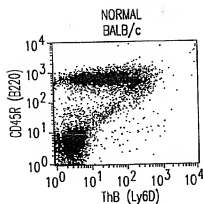


FIG. 11B

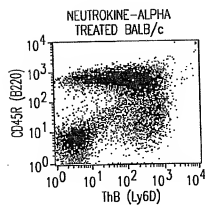


FIG. 11C

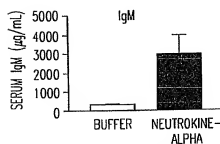


FIG. 11D

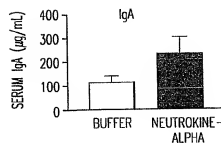


FIG. 11E

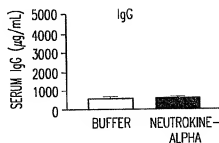


FIG. 11F